Inter-Agency Conference on Metabolic Engineering 2005

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Introduction of Prof. John W. Frost

by

Dr. Harold J. Bright
ONR Molecular Biomimetics Program



Green Synthesis: Connecting DNA Sequence Space with Chemical Manufacture

MEDICAL AND BIOLOGICAL S&T DIVISION





Propelling Research

By MEGHAN GILBERT

The State News

A team of three MSU scientists has found a way to use bacteria in order to make the chemical used to propel Navy Hellfire missiles.



Chemistry professor **John Frost**, who grew up in Illinois around the time Penicillin was discovered, always has been interested in biosynthesis, the biological way to combine parts of elements to form a whole.





The Portfolio

Renewable Polyols Engineered Microbes

Shikimic Acid (Tamiflu)

Phloroglucinol ONR (EM precursor for Navy/MC)

Butanetriol ONR/NSF (EM precursor for Navy/MC)

Caprolactam (Nylon)



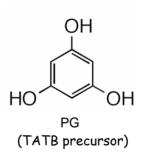
Rationale for Navy/MC Green Synthesis of Energetic Materials

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State of the EM Art

Biocatalytically Impossible!

Realistic Targets



Bionitration

<u>Issues</u>

- Economics of EM manufacture
- Feedstock security BT and PG are now available only in China/India
- Coordination with ATK-Thickol on PG → TATB
- Strong dual-use is essential to get industry involved in bioprocessing (e.g. chiral synthons in BT biosynthesis are cholesterol drug intermediates; PG → catechol, a high volume coating)



Green Synthesis of Navy/MC Energetic Compounds: Butanetriol (BT) as a Case Study

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J. Frost, Michigan State U.

Also NSWCIH, Michigan Biotech Inst.

ONO2

O2NO

ONO2

<u> 811N</u>:

BT

BTTN

- · made from BT (now from China!)
- · used in propellants (e.g., Hellfire, 15K lb/yr)
- better than nitroglycerin (NG, >5M lb/yr)
- · hasn't replaced NG because BT is ~ \$40/lb owing to dirty BT synthesis

Engineer Cost-Effective Microbial BT Synthesis

- · no metals, organics, salt streams
- · no high temperatures or pressures
- · single, engineered microbe
- · renewable biofeedstocks ($CO_2 \rightarrow sugars$)



Bottom Line:

microbial synthesis of BT at ≤ \$10/lb that meets military specs

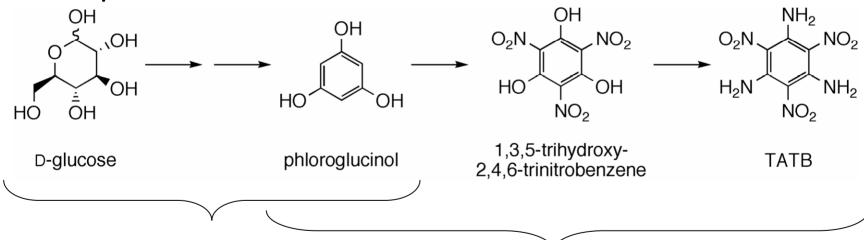


Phloroglucinol > TATB

MEDICAL AND BIOLOGICAL S&T DIVISION MURI

J. Frost, Michigan State U. Also Integrated Genomics, U. Illinois, Rice U.

- Microbial production of PG from glucose at 1g/liter
- · This permits collaboration with ATK-Thiokol



ONR/DARPA MURI (microbial)

ATK-Thiokol (chemical)

· TATB will then be from domestic source